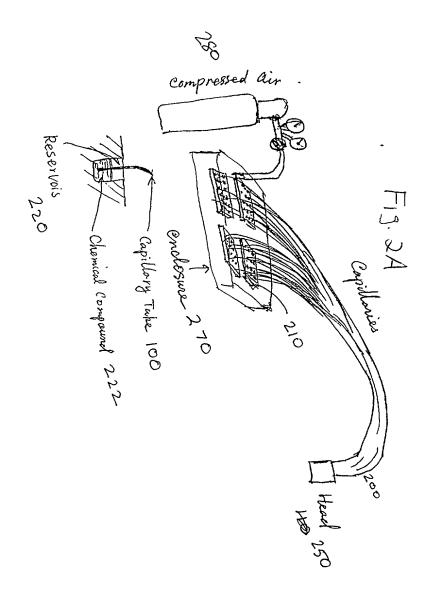
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Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES
TOR HIGH THROUGHPUT SCREENING
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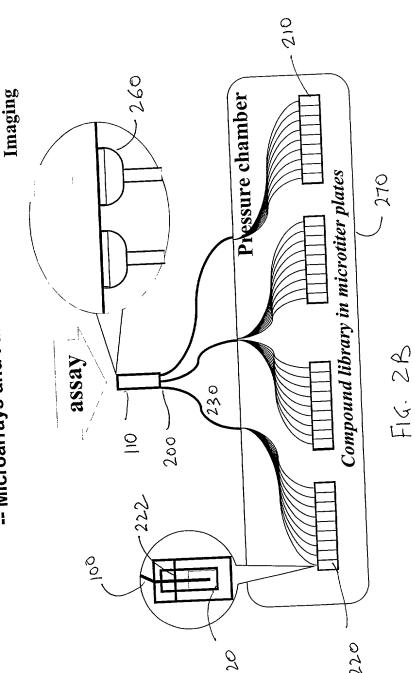
Figure 1



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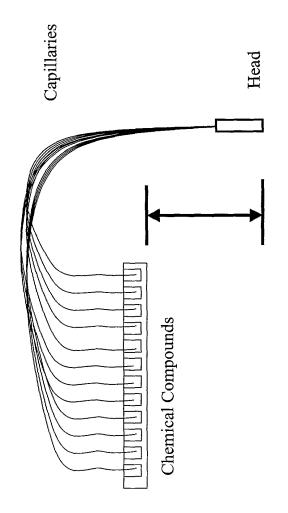
Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES
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-- Microarrays and Fiber Bundles



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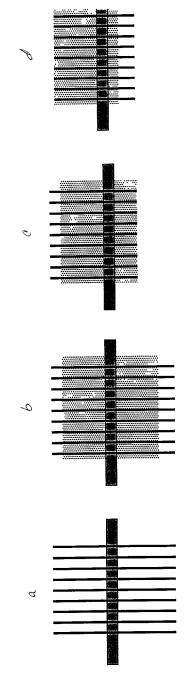


Fig. 4. Fabrication of delivery head using a guide plate

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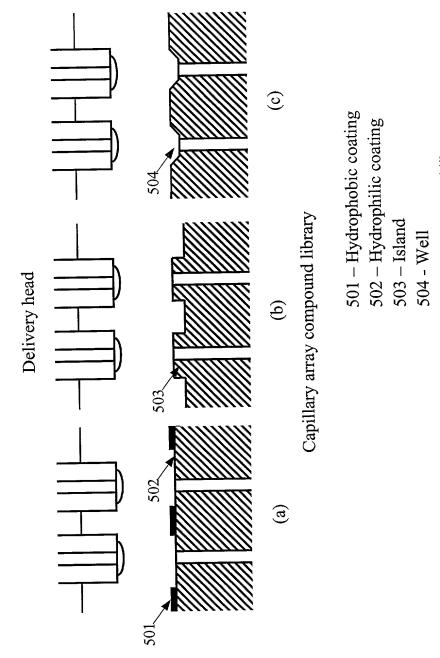


Fig. 5. Surface features on the surface of the capillary array compound library to prevent crosscontamination during compound loading

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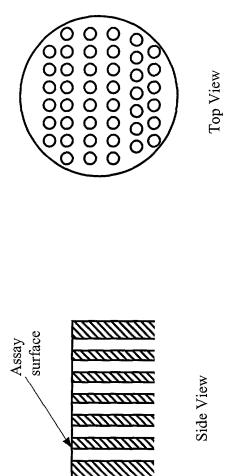


Fig. 6. Basic configuration of capillary array substrate for the portable compound library

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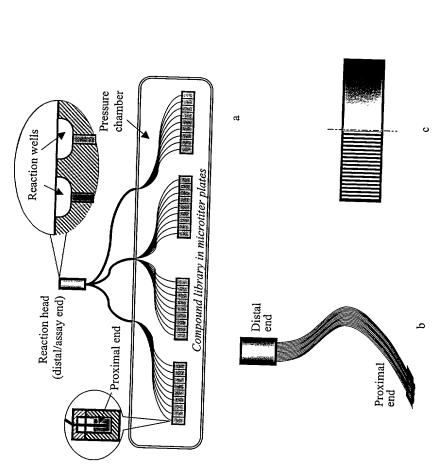
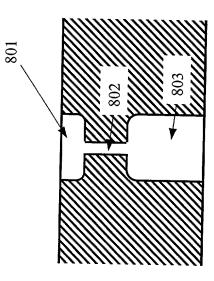


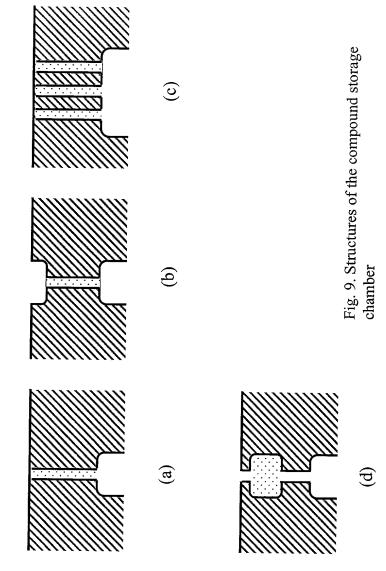
Fig. 7 The capillary array compound library in different formats



801 – Mixing/reaction well 802 – Flow regulator for reagent metering 803 – Compound reservoir

Fig. 8. Internal structure of a through hole in capillary array compound library

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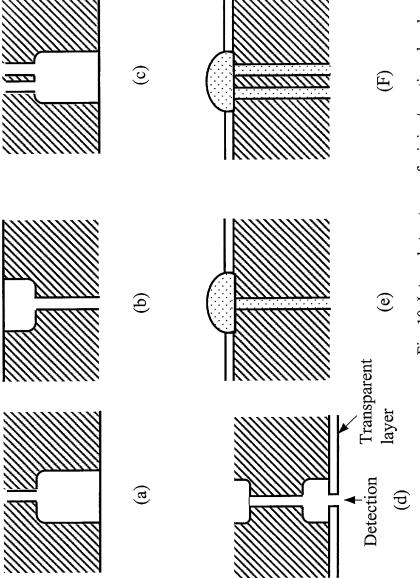


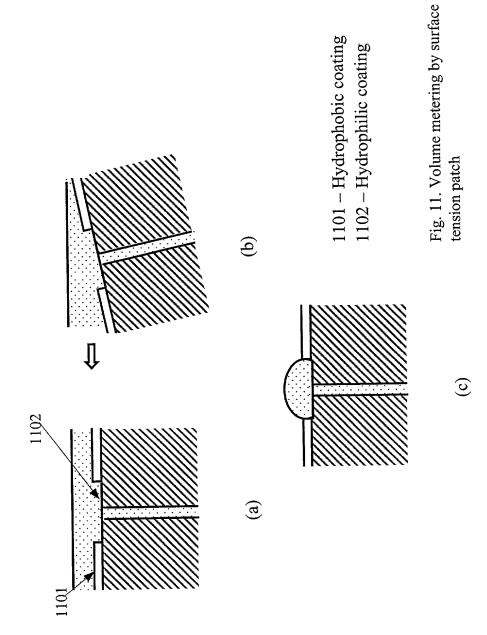
Fig. 10. Internal structures of mixing/reaction chamber

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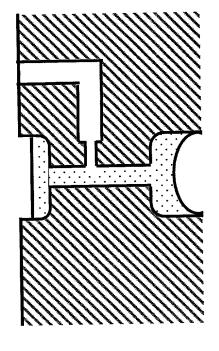
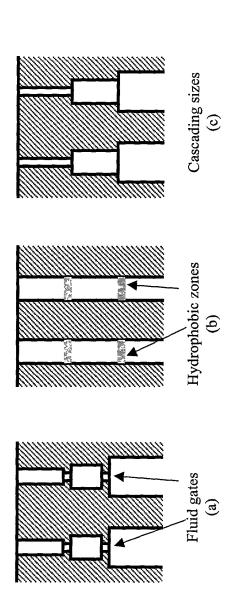


Fig. 12. Fluid regulator with side air tunnel

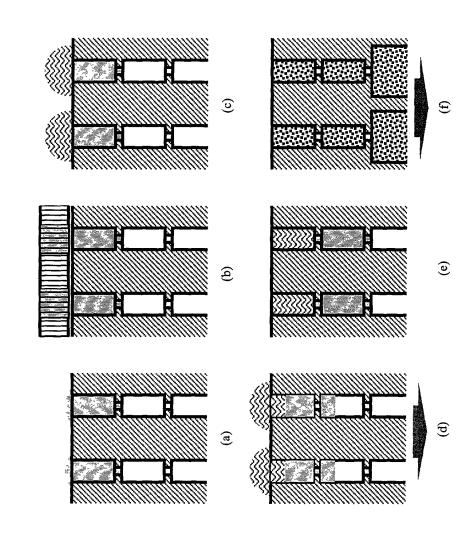
Fig. 13 Internal through hole structures to facilitate chamber volume metering and mixing



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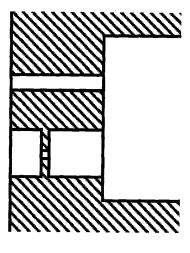
Fig. 14 Process of metering multiple reagents using interconnected chambers



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Fig. 15 Special through hole structure where multiple chambers links to a chamber in parallel



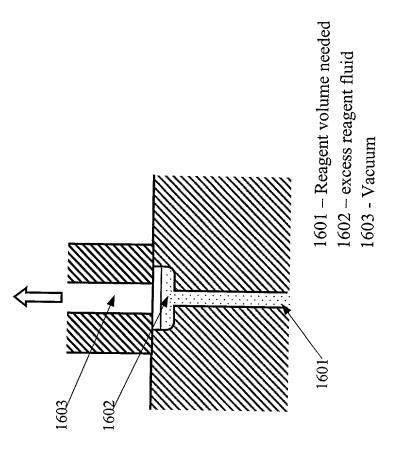


Fig.16. Removal of excess fluid by vacuum

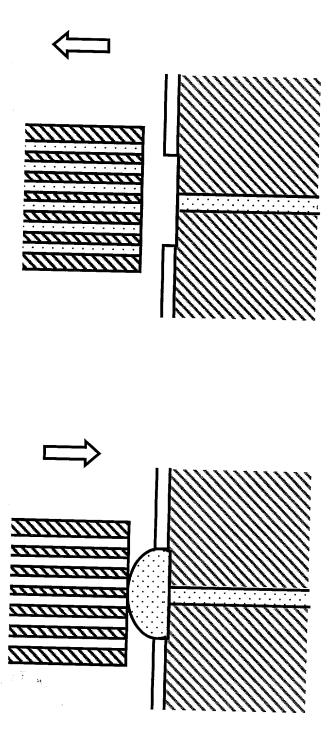


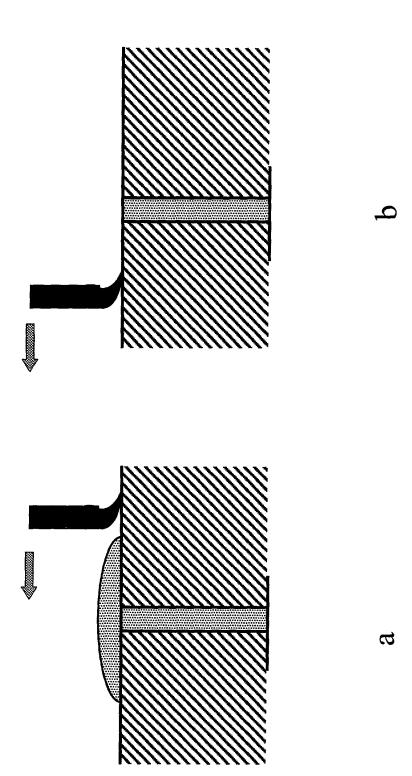
Fig. 17. Excess fluid removal using a second capillary array

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Fig. 18. Excess Fluid Removal by Wiping



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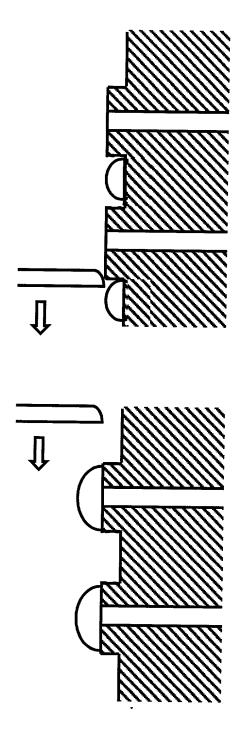
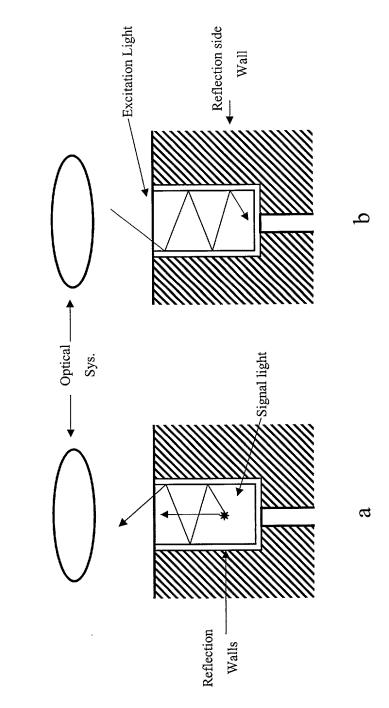


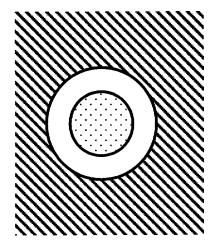
Fig.19. A method for reducing cross-contamination between adjacent holes during excess fluid removal

Fig. 20. Using Reflection Wall of Reaction Chamber to Enhance Optical Signal of the Assay



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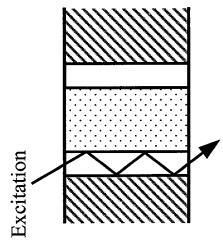


Fig 21. Light guiding capillary

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1. Immobilization anglass

EtQ H
EtCST H
EtQ C
EtQ E
EtQ C
E

n=0, 1, 8 R=-CH₃ -C₂H₃ -C₂H₄-

FIG 22A

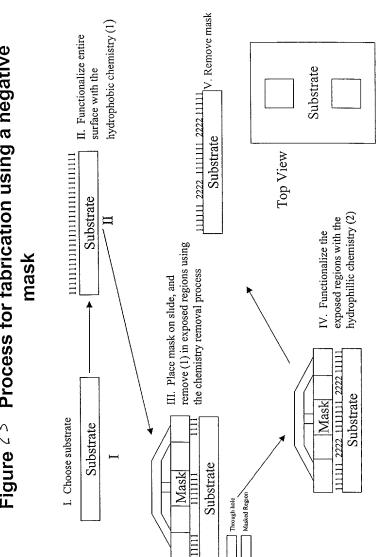
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-NO₂ (EtO)₃SiH Si(OEt) 3 solution phase silanization O₂N O₂N´ UV-light 350nm O₂N O₂N O₂N

FIG ZZC

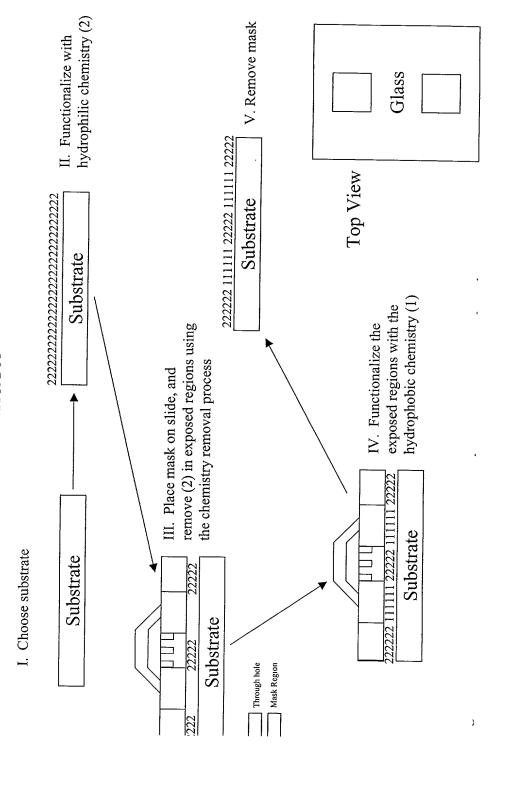
Process for fabrication using a negative Figure 23



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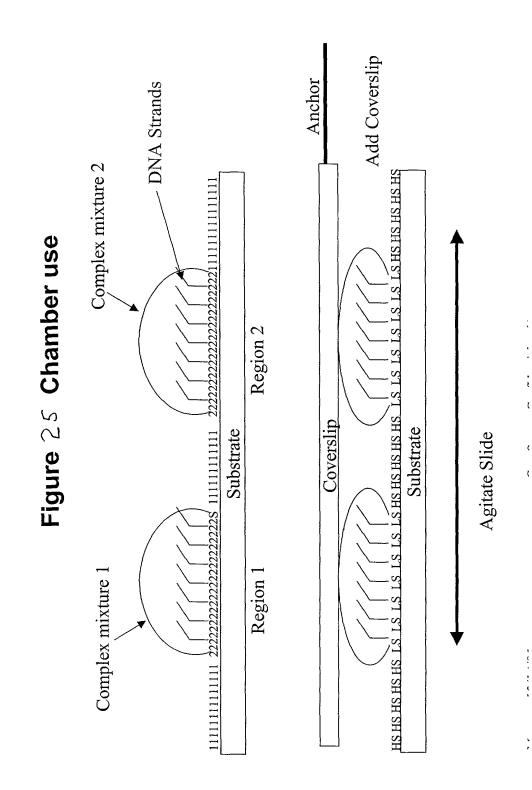
Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES
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Figure 24 Process for the fabrication using positive mask



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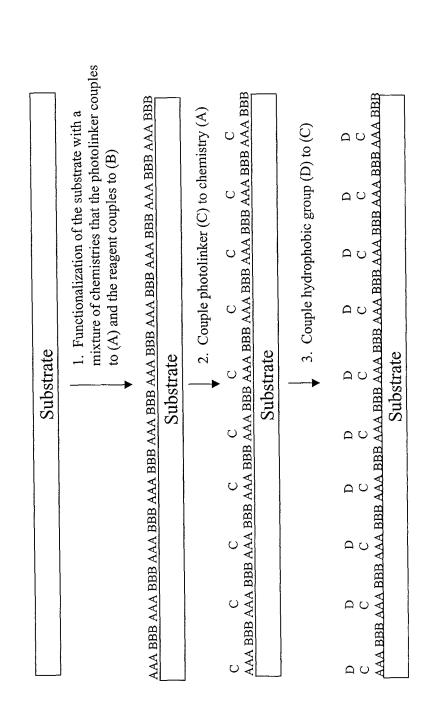
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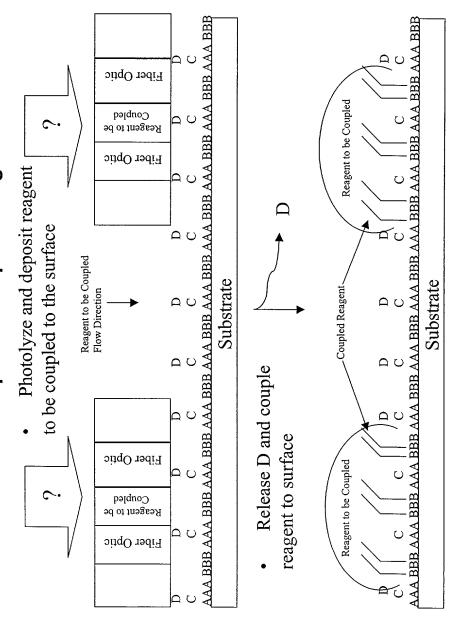
FOR HIGH THROUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: 10/080,274 Docket No.: 473532000620

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Figure 264 Surface Tension Patterning: On-capillary Fiber optic based patterning



Surface Tension Patterning: On-capillary Fiber optic based patterning Figure 2 6B



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1050E74 O7150E

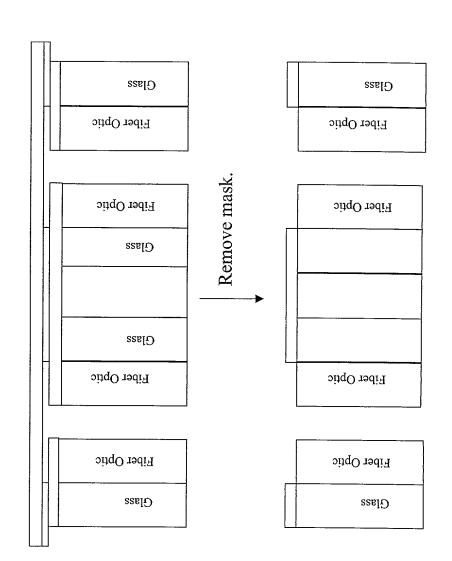
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Coat Surface with a Hydrophobic Reagent Volume Metering using Surface Glass Glass Fiber Optic Fiber Optic **Tension Features** Fiber Optic Fiber Optic Glass Glass Glass Glass Fiber Optic Fiber Optic **Fiber Optic** Fiber Optic Glass GJsss

Figure 27B Volume Metering using Surface Tension Features

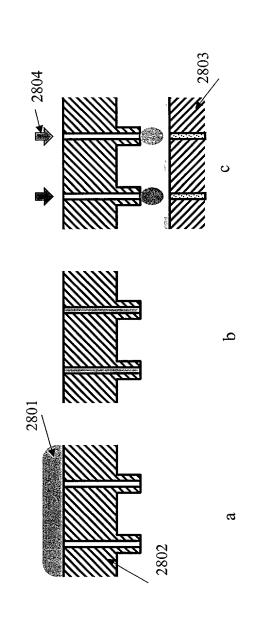
Place a Mask on to the Surface and Expose the Surface to the Chemistry Removal Process



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Fig. 28 Reagent pre-metering using an intermediary through-hole array



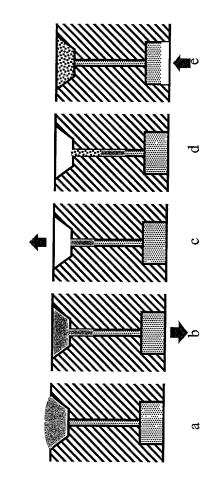
2801 - reagent fluid applied in excessive;

2804 - pressure

^{2802 -} intermediary through hole array;

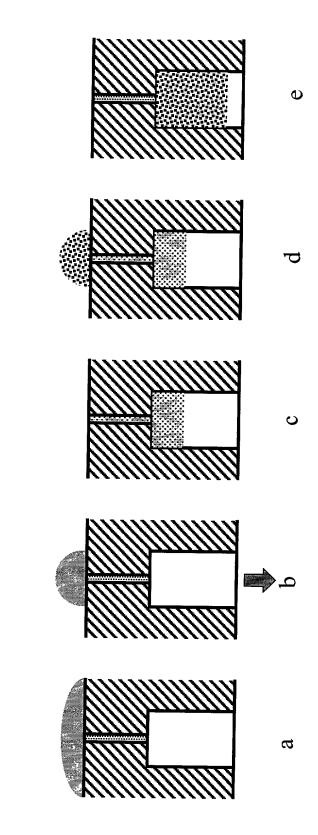
^{2803 -} capillary array compound library;

Fig. 29 Metering and mixing with a multi-use capillary array compound library



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Fig. 30 Metering with hydrophilic patch and mixing

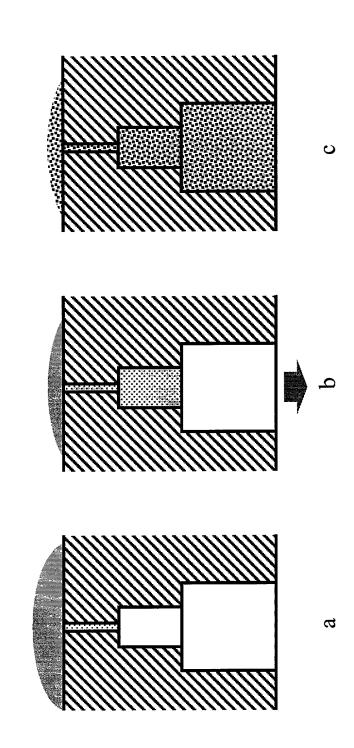


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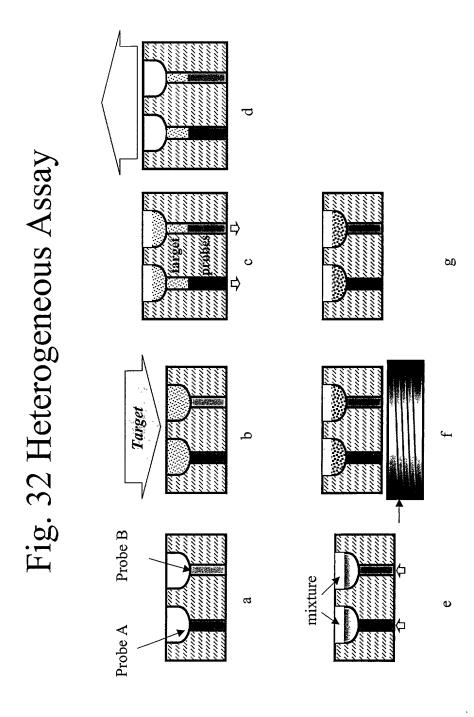
Fig. 31 Mixing and metering with interconnected chambers



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Antibody Immobilization via the Carbohydrate Moiety

1. Oxidation of antibodies vicinal diol group to its aldehyde

2. Conjugation of maleimide moiety with antibody

3. Immobilization of the modified antibody to the surface.

Immobilization via Amine Goups

1. Hydrosilylation of (3-mercaptopropyl)triethoxysilane on the surface of fiber

2. Formation of a thioether bond

3. Attachment of fiber to antibody

Antibody Immobilization via Streptavidin

1. Label antibody with biotin

2. Modification of fiber surface with biotin maleimide

Meeting 12/14/01

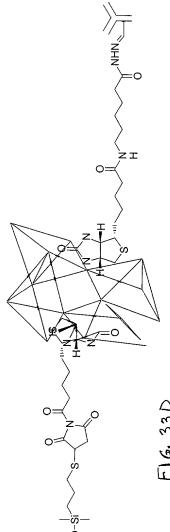
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Antibody Immobilization via Streptavidin

3. Conjugate Streptavidin to the surface

4. Conjugate Biotin Anitbody to the surface



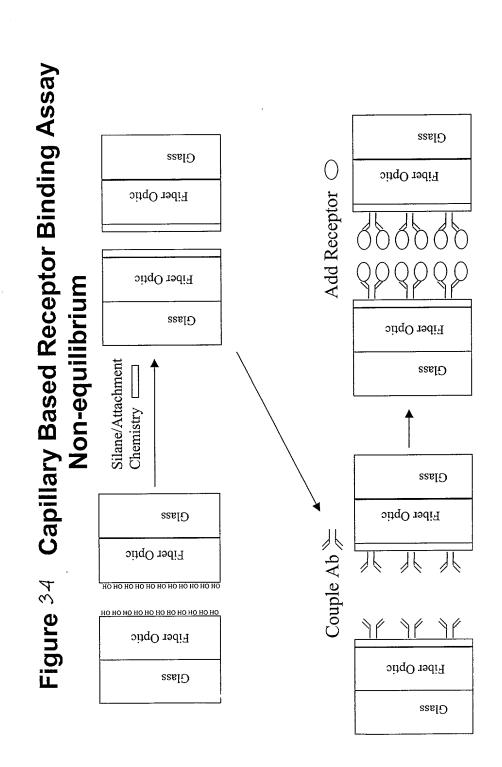
F16.33D

Formation of thiazolidine

1. Surface attachment and formation of the linker

2. Thiazolidine formation

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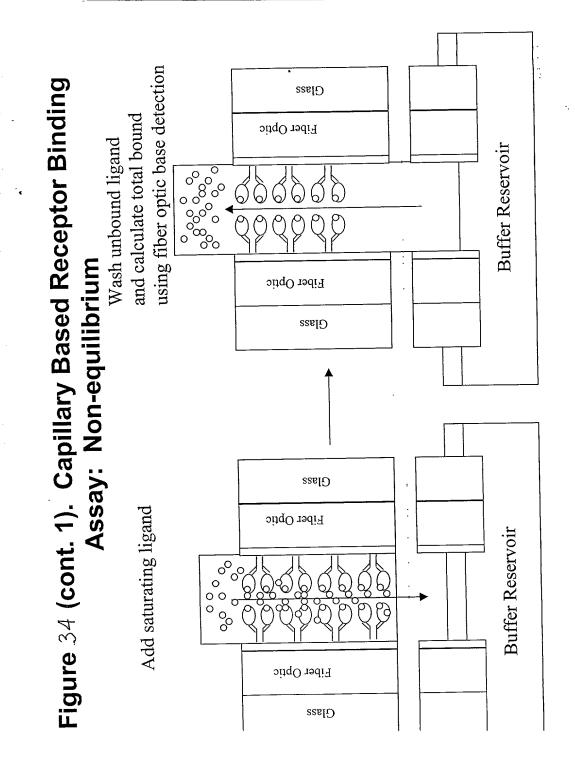


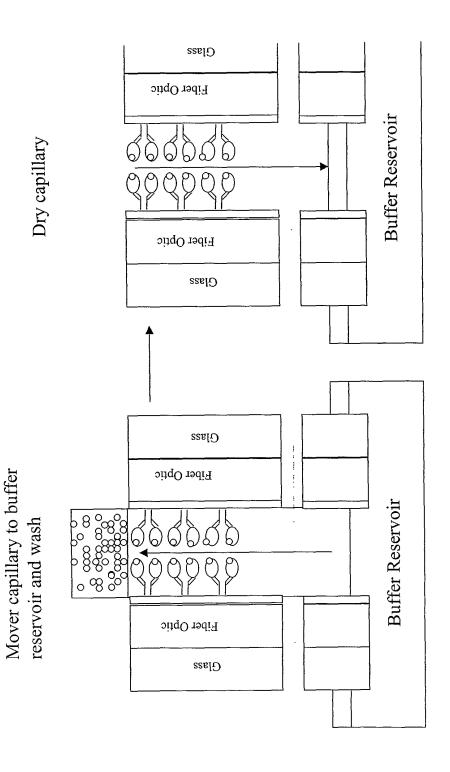
Figure 34 (cont. 2). Capillary Based Receptor Binding Assay: Non-equilibrium

0 °° 8 0 optic based detection to observe Add compound and use fiber 0 Glass 0 0 Fiber Optic 00088000 0000000 08000000 0 kinetics Fiber Optic 0 0 Glass 000 0 00 00 0 00 8 0 Secondage Servor Glass Fiber Optic compound reservoir Move capillary to Fiber Optic 0 ∞ Glass $b \infty$ 0 80 0 0 00

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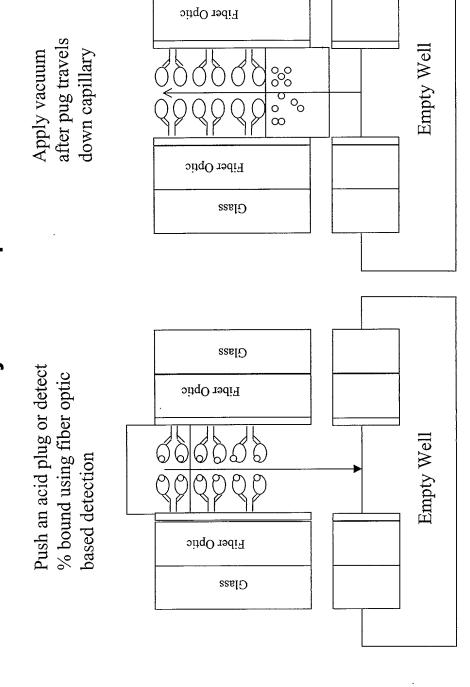
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Figure 34 (cont. 3). Capillary Based Receptor Binding Assay: Non-equilibrium



Glass

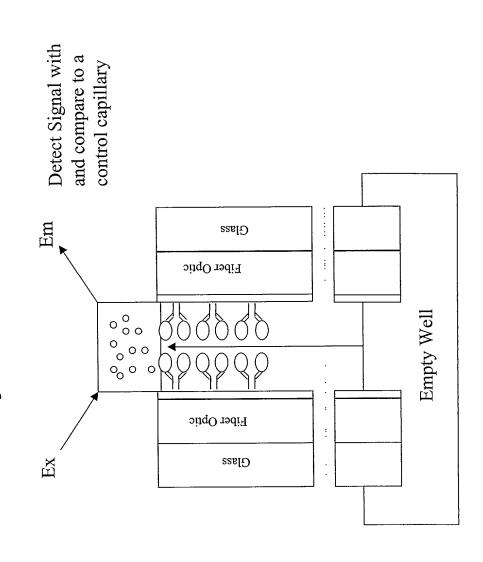
Figure 34 (cont. 4). Capillary Based Receptor Binding Assay: Non-equilibrium



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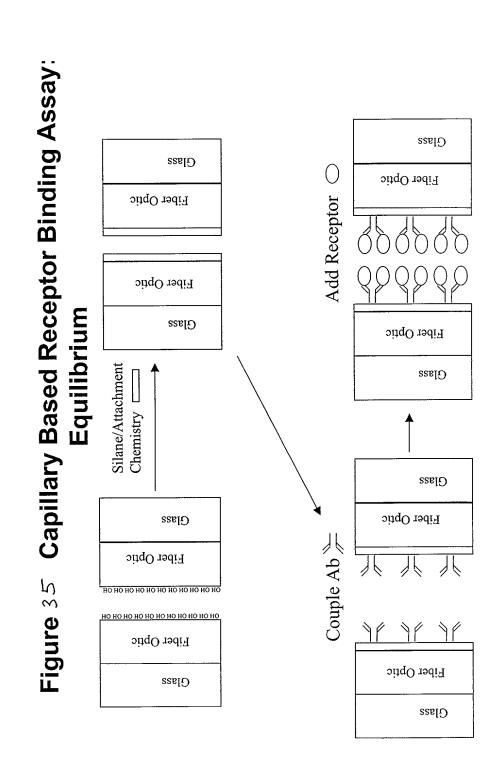


Figure 35 (cont. 1). Capillary Based Receptor Binding Assay: Equilibrium

compound/ligand reservoir. Move Capillary to

Add solution and let system reach equilibrium. Detect equilibrium using fiber optic base detection.

0 0 Glass Compound/LigandReservoir Fiber Optic 88, 00000 0 00 0 bo 0 0 Fiber Optic Glass 0 0 Glass Fiber Optic Fiber Optic Glass ∞ 8% °8°

Figure 35 (cont. 2). Capillary Based Receptor Binding

Class Fiber Optic **Buffer Reservoir** Dry Capillary Fiber Optic Assay: Equilibrium Class reservoir and wash capillary with Glass buffer. Detect % bound using fiber optic based detection. Move capillary to a buffer Fiber Optic **Buffer Reservoir** Fiber Optic Glass

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after pug travels

Apply vacuum

down capillary

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Glass Detect signal using fiber optic base detection or elute bound Fiber Optic ligand with acid. Fiber Optic

Glass Fiber Optic Empty Well 000 0 Fiber Optic Glass Empty Well Glass

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Figure 35 (cont . 4). Capillary Based Receptor Binding Assay: Equilibrium

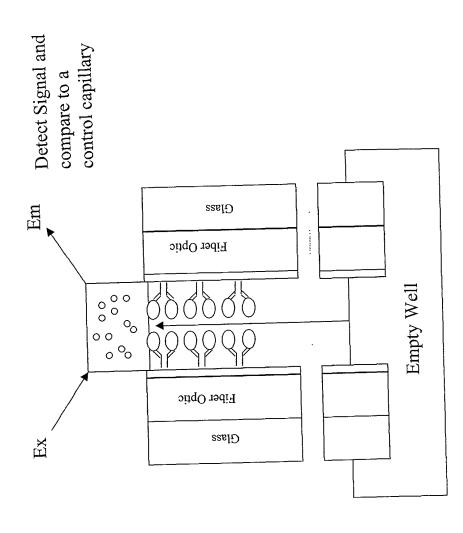
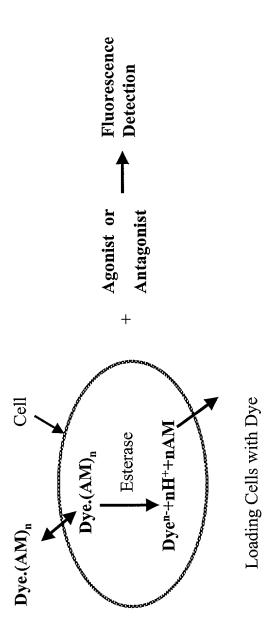


Figure 36



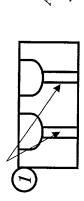
Assay Based on Tracking Cytosolic [Ca++]

Figure 37

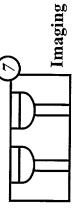
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Protein Array & Cell Array

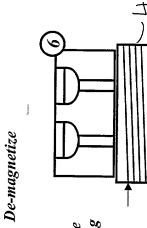
Library of antigen or antibody Attached to magnetic beads



Aspiration from top



Mixing circle



circle



Seal,

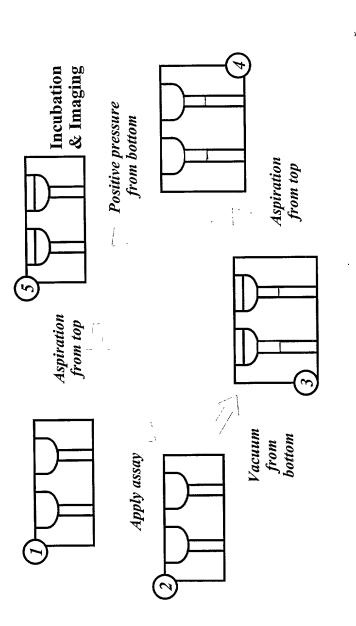
Binding interaction

38A Fla.

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FIG. 38B

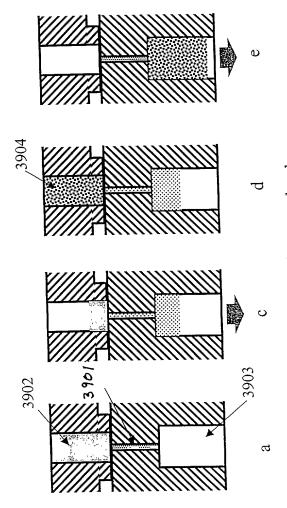
Fluid-Array™ XHTS



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Fig. ψ_0 Metering with through hole plates and mixing



3902 - reagent A (i.e. enzyme) in through hole plate A 3901 - compound and compound storage chamber

3903 - mixing/reaction chamber

3904 - reagent B (i.e. substrate) in through hole plate B

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Fig. 39 One embodiment of the capillary array cartridge design

